

<120> PRODUCTION OF PHARMACEUTICAL PROTEINS IN TRANSGENIC PLASTIDS

<140> 09/807,742

<141> 2001-04-18

<150> PCT/US01/06288

<151> 2001-02-28

<160> 19

<170> PatentIn Ver. 2.1

<210> 1

<211> 1250

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Synthetic peptide

<220>

<223> This sequence may encompass 1-250 Gly Val Gly Val Pro repeats

<400> 1

Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly
1 5 10 15

Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val
			20					25					30		

Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly
35 40 45

Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val
50 55 60

Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro
65					70					75					80

Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly
85 90 95

Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val
			100					105					110		

Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly
115 120 125

Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val
130						135					140				
Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro
145					150					155					160
Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly
				165					170					175	
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val
			180					185					190		
Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly
		195					200					205			
Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val
		210				215					220				
Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro
225					230					235					240
Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly
				245					250					255	
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val
			260					265					270		
Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly
		275					280					285			
Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val
		290				295					300				
Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro
305					310					315					320
Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly
				325					330					335	
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val
			340					345					350		
Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly
		355					360					365			
Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val
		370				375					380				
Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro
385					390					395					400
Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly
				405					410					415	
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val
			420					425					430		

Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Pro Gly	Val Gly	Val Pro	Gly Val	Pro Gly	Val Gly	Val Gly
	435				440				445			
Val Pro	Gly Val	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Gly	Val Gly
	450				455				460			
Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Pro Gly
	465				470				475			480
Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Gly Val	Pro Gly	Val Gly	Pro Gly
			485					490				495
Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Val Gly	Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Gly
		500				505				510		
Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Pro Gly	Val Gly	Gly Val	Pro Gly	Val Gly	Val Gly	Gly Val
	515					520				525		
Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val	Val Gly
	530					535				540		
Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Val Gly	Pro Gly
	545					550			555			560
Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Pro Gly	Val Gly	Gly Val	Pro Gly	Gly Val
			565					570				575
Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Val Gly	Pro Gly	Val Gly	Gly Val	Pro Gly	Gly Val	Val Gly
			580					585			590	
Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Pro Gly	Val Gly	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val
	595					600				605		
Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val	Val Gly
	610					615				620		
Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Val Gly	Pro Gly
	625					630			635			640
Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Pro Gly	Gly Val
			645					650				655
Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Val Gly	Pro Gly	Val Gly	Gly Val	Pro Gly	Gly Val	Val Gly
			660					665			670	
Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Pro Gly	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val
	675						680			685		
Val Pro	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Gly Val	Val Gly
	690					695				700		
Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Gly Val	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val	Val Gly	Pro Gly
	705					710			715			720
Gly Val	Gly Val	Pro Gly	Val Gly	Val Gly	Val Pro	Gly Val	Pro Gly	Gly Val	Gly Val	Pro Gly	Gly Val	Gly Val
			725					730				735

Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val
 740 745 750
 Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly
 755 760 765
 Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val
 770 775 780
 Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro
 785 790 795 800
 Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly
 805 810 815
 Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val
 820 825 830
 Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly
 835 840 845
 Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val
 850 855 860
 Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro
 865 870 875 880
 Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly
 885 890 895
 Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val
 900 905 910
 Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly
 915 920 925
 Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val
 930 935 940
 Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro
 945 950 955 960
 Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly
 965 970 975
 Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val
 980 985 990
 Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly
 995 1000 1005
 Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val
 1010 1015 1020
 Pro Gly Val Gly Val Pro Gly Val Gly Val Pro Gly Val Gly Val Pro
 1025 1030 1035 1040

5

Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly				
1045					1050					1055									
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val				
1060					1065					1070									
Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly				
1075					1080					1085									
Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val				
1090					1095					1100									
Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro				
1105					1110					1115					1120				
Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly				
1125					1130					1135									
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val				
1140					1145					1150									
Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly				
1155					1160					1165									
Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val				
1170					1175					1180									
Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro				
1185					1190					1195					1200				
Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly				
1205					1210					1215									
Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val				
1220					1225					1230									
Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Val	Gly				
1235					1240					1245									
Val	Pro																		
1250																			

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<210> 2
<211> 6
<212> PRT
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Illustrative
endoplasmic reticulum retention signal

<400> 2
Ser Glu Lys Asp Glu Leu
1 5

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<210> 3
<211> 4
<212> PRT
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Illustrative peptide

<400> 3
Gly Pro Gly Pro
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<210> 4
<211> 25
<212> DNA
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Primer

<400> 4
ccgtcgacgt agagaaqtcc gtatt 25

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<210> 5
<211> 27
<212> DNA
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Primer

<400> 5
gccccatggtta aaatcttgggt ttatttta 27

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<210> 6
<211> 28
<212> DNA
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Primer

<400> 6
cctttaaaaa gccttcatt ttctattt 28

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<210> 7
<211> 25
<212> DNA
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Primer

<400> 7
gccatggtaa aatcttggtt tatta

25

<210> 8
<211> 12
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Illustrative
preferred nucleotide sequence

<400> 8
tttcgtttca gt

12

<210> 9
<211> 5
<212> PRT
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Synthetic
peptide

<400> 9
Ala Val Gly Val Pro
1 5

<210> 10
<211> 7
<212> PRT
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Illustrative
peptide

<400> 10
Glu Asn Leu Tyr Phe Gln Gly
1 5

<210> 11
<211> 6
<212> PRT
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Illustrative
peptide

<400> 11
Leu Val Pro Arg Gly Ser
1 5

<210> 12
 <211> 6
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Description of Artificial Sequence: 6-His tag

<400> 12
 His His His His His His
 1 5

<210> 13
 <211> 25
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Description of Artificial Sequence: Primer

<400> 13
 aaaaccgctc ctcagttcgg attgc 25

<210> 14
 <211> 25
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Description of Artificial Sequence: Primer

<400> 14
 ccgcgttggt tcatcaagcc ttacg 25

<210> 15
 <211> 119
 <212> PRT
 <213> Escherichia coli

<400> 15
 Gly Ile Val Pro Gly Val Gly Ile Val Pro Gly Val Gly Ile Val Pro
 1 5 10 15
 Gly Val Gly Ile Val Pro Gly Val Gly Ile Val Pro Gly Val Gly Ile
 20 25 30
 Val Pro Gly Val Gly Ile Val Pro Gly Val Gly Ile Val Pro Gly Val
 35 40 45
 Gly Ile Val Pro Gly Val Gly Ile Val Pro Gly Val Gly Ile Val Pro
 50 55 60
 Gly Val Gly Ile Val Pro Gly Val Gly Ile Val Pro Gly Val Gly Ile
 65 70 75 80

Gly Val Gly Val Pro Gly Val
115

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<210> 16
<211> 260
<212> DNA
<213> Homo sapiens
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<400>	16						
tttgtgaacc	aacacctgtg	cggtcacac	ctggtggaag	ctctctacct	agtgtgcggg	60	
gaacgaggct	tcttctacac	acccaagacc	cgccgggagg	cagaggacct	gcaggtgggg	120	
caggtggagc	tgggcggggg	ccttgggtgca	ggcagcctgc	agcccttggc	cctggagggg	180	
tccttcgaga	agcgtggcat	tgtggaacaa	tgctgtacca	gcattctgctc	cctctaccag	240	
ctggagaact	actgcaacta					260	

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<210> 17
<211> 260
<212> DNA
<213> Artificial Sequence
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<220>
<223> Description of Artificial Sequence: Chloroplast modified proinsulin sequence

<400> 17						
ttcgtaaacc	aacacttatg	tggttctcac	ctagtagaag	ctttatactt	agtatgtggt	60
gaacgtgggt	tcttctacac	tcctaaaact	cgtcgtgaag	ctgaagattt	acaagtaggt	120
caagtagaat	taggtgggtg	tcttgggtgt	ggttctttac	aaccttttagc	tttagaaggt	180
cttttacaaa	aacgtggtat	tgtagaacaa	tgttgtaact	ctatttggtc	tttataccaa	240
ttagaaaact	actgtaacta					260

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<210> 18
<211> 210
<212> DNA
<213> Homo sapiens
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```
<400> 18
ggaccggaga cgctctgcgg ggctgagctg gtggatgctc ttcagttcgt gtgtggagac 60
aggggctttt atttcaacaa gcccacaggg tatggctcca gcagtcggag ggcgcctcag 120
acaggcatct tggatgagtg ctgtctccgg agctgtgatt taaggaggct ggagatgtat 180
tcgcaccccc tcaagctctg caagtgcagt
210
```

```
<210> 19
<211> 210
<212> DNA
<213> Homo sapiens
```

<400> 19

ggtcctgaaa	ctttatgtgg	tgctgaatta	gtagatgctt	tacaattcgt	atgtggatgat	60
cgtggtttct	atttcaacaa	acctactggg	tacggttctt	cttctcgctg	tgctcctcaa	120
actggatttg	tagatgaatg	ttgtttccgt	tcttgatgatt	tacgtcggtt	agaaatgtac	180
tggtgctcct	taaaacctgc	taaatctgct				210